```
related urticaria), hypertensive toxemia, glomerular
endotheliosis and
CC
     cholestasis. The present sequence represents a homolog of the
human
CC
    pregnancy zone protein precursor-like protein NOV10. Note:
The sequence
     data for this patent is also available in electronic format
directly from
CC
     the US patent office at
CC
     seqdata.uspto.gov/sequence.html?DocID=20060009634.
XX
SQ
     Sequence 1450 AA;
  Query Match
                        100.0%; Score 88; DB 10; Length 1450;
 Best Local Similarity 100.0%; Pred. No. 9.5e-06;
  Matches
           18; Conservative 0; Mismatches
                                                  0;
                                                      Indels
0; Gaps
            0;
Qy
            1 LLIYAVLPTGDVIGDSAK 18
              Db
          516 LLIYAVLPTGDVIGDSAK 533
RESULT 18
AAU81018
     AAU81018 standard; protein; 1451 AA.
ID
XX
AC
    AAU81018;
XX
DT
    09-APR-2002 (first entry)
XX
DE
     Human alpha2 macroglobulin (alpha2M) receptor #1 mature
protein.
XX
KW
     Human; mouse; alpha2 macroglobulin; receptor; alpha2M; HSP;
KW
     heat shock protein; alpha2M receptor-HSP complex; autoimmune
disorder;
KW
    multiple sclerosis; rheumatoid arthritis; endocytosis;
inflammation;
     cytokine clearance; antigen presentation disruption;
carcinoma; sarcoma;
    proliferative disorder; cancer; infectious disease; bacterial
     intracellular parasite; hypercholesterolaemia; protozoan
infection;
    Alzheimer's disease; diabetes; osteoporosis; viral infection;
```

```
protein.
XX
OS
     Homo sapiens.
XX
PN
     WO200192474-A1.
XX
PD
     06-DEC-2001.
XX
PF
     04-JUN-2001; 2001WO-US018041.
XX
PR
     02-JUN-2000; 2000US-0209095P.
     25-JUL-2000; 2000US-00625137.
PR
PR
     22-SEP-2000; 2000US-00668724.
PR
     28-DEC-2000; 2000US-00750972.
XX
PA
     (UYCO-) UNIV CONNECTICUT HEALTH CENT.
XX
PI ·
     Srivastava PK;
XX.
DR
     WPI; 2002-122061/16.
XX
PT
     Screening assays for identifying compounds useful for
treating immune
     disorders, comprises identification of compounds that
modulate alpha 2-
PT
     macroglobulin receptor-heat shock protein interaction.
XX
PS
     Disclosure; Fig 13B; 236pp; English.
XX
CC
    The invention relates to screening assays comprising
identification of
CC
     compounds that modulate alpha2 macroglobulin (alpha2M)
receptor (which
CC
     also functions as a heat shock protein (HSP) receptor)-HSP
interaction. A
     compound that modulates the activity of an alpha2M
receptor-HSP complex
CC
     can be identified by contacting the compound with HSP and
alpha2M
CC
     receptor and measuring the level of alpha2M activity or
expression. If
CC
     the level differs from that perceived in the absence of the
test
CC
     compound, a compound that modulates an alpha2M
receptor-HSP-mediated
     process is identified. The identified compounds are useful
```

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for treating
     autoimmune disorders (such as multiple sclerosis or
CC
rheumatoid
     arthritis), diseases or disorders involving disruption of
antigen
    presentation, endocytosis, cytokine clearance or
CC
CC
    proliferative disorders (such as cancers including sarcomas
and
CC
     carcinomas), infectious diseases (such as those caused by
viruses,
    bacteria, protozoans and intracellular parasites),
hypercholesterolaemia,
    Alzheimer's disease, diabetes and osteoporosis. Sequences
CC
AAU81016-
     AAU81073 represent human and mouse alpha2M receptors and
peptide
CC fragments of the invention
XX
SQ
     Sequence 1451 AA;
                         100.0%; Score 88; DB 5; Length 1451;
 Query Match
 Best Local Similarity 100.0%; Pred. No. 9.5e-06;
 Matches
           18; Conservative 0; Mismatches 0; Indels
0; Gaps
           0;
           1 LLIYAVLPTGDVIGDSAK 18
Qу
             Db
         517 LLIYAVLPTGDVIGDSAK 534
RESULT 19
ADK41537
ΙD
    ADK41537 standard; protein; 1451 AA.
XX
AC
    ADK41537;
XX
DT
     06-MAY-2004 (first entry)
XX
DΕ
    Anti-cell surface antigen related protein #21.
XX
     cytostatic; immunosuppressive; gene therapy; anti-cell
surface antigen;
     CD84Hyl; alpha2MHy; IgFBP-7Hyl; Toll-like receptor 9; VpreB1;
antibody;
     lymphoma; cancer; autoimmune disorder; systemic lupus
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d his

(FILE 'HOME' ENTERED AT 09:54:13 ON 15 AUG 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 09:54:53 ON 15 AUG 2007

| | AUG | 2007 | | | | | | | | | |
|-----|-----|-------|----|-----------|------|----|--------|------|------|------------|----------|
| L1 | | 21001 | S | (AĹ | PHA | 2 | MACRO | GLO | BUL: | IN) | |
| L2 | | 363 | S | L1 | AND | DI | ABETE | ? | | | |
| L3 | | 20 | S | L2 | AND | UR | INE? | | | | |
| L4 | | 14 | DI | UPLI | CATE | R | REMOVE | : L3 | (6 | DUPLICATES | REMOVED) |
| L5 | | 11 | S | L4 | AND | PD | <2004 | | | | |
| L6 | | 0 | S | L2 | AND | (M | IASS S | PEC |) | | |
| L7 | | 21 | S | L2 | AND | MA | SS | | | | |
| T8 | | 13 | DI | UPLI | CATE | R | REMOVE | L7 | (8 | DUPLICATES | REMOVED) |
| L9 | | 0 | S | $\Gamma8$ | AND | PD | ,2004 | | | | |
| L10 | • | 4 | S | T8 | AND | PD | <2004 | ı | | | |

(FILE 'HOME' ENTERED AT 09:54:13 ON 15 AUG 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 09:54:53 ON 15 AUG 2007

| | 1700 | 2007 | | |
|------|------|-------|--------------------------------------------|---|
| L1 | | 21001 | S (ALPHA 2 MACROGLOBULIN) | |
| L2 | | 363 | S L1 AND DIABETE? | |
| L3 | | 20 | S L2 AND URINE? | |
| L4 | | 14 | DUPLICATE REMOVE L3 (6 DUPLICATES REMOVED) | |
| L5 | | 11 | S L4 AND PD<2004 | |
| L6 | | 0 | .S L2 AND (MASS SPEC) | |
| L7 | | 21 | S L2 AND MASS | • |
| L8 | | 13 | DUPLICATE REMOVE L7 (8 DUPLICATES REMOVED) | |
| L9 . | | 0 | S L8 AND PD, 2004 | |
| L10 | | 4 | S L8 AND PD<2004 | |

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```
ANSWER 2 OF 11 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
     1995:509170 BIOSIS
ΑN
     PREV199598514220
DN
ΤI
     A radioimmunometric assay for urinary alpha-2-
     macroglobulin.
ΑU
     Ito, Seiki; Usami, Akio; Yamazaki, Masatoshi; Shibata, Akira
     Div. Gerontol., Akita Univ. Hosp., Akita 010, Japan
CS
SO
     Tohoku Journal of Experimental Medicine, (1995) Vol. 176, No. 3,
     pp. 137-147.
     CODEN: TJEMAO. ISSN: 0040-8727.
DT
     Article
LA
     English
ED ·
     Entered STN: 29 Nov 1995
     Last Updated on STN: 29 Nov 1995
     To measure urinary alpha-2-macroglobulin
     levels, a sensitive radioimmunometric assay was established. The least
     detectable level of this assay was 225 pg/ml. A linear correlation
     between alpha-2-macroglobulin levels and
     serial dilution of urine samples was found. Western blot
     analysis and study on column chromatography revealed that the molecular
     weight of alpha-2-macroglobulin in
     urine was identical to that of serum alpha-2-
     macroglobulin. The findings suggested that urinary substance
     detected by the present assay was truly alpha-2-
     macroglobulin. Timed overnight urine samples from 49
     diabetic patients with retinopathy and 20 healthy controls were measured
     by the present assay. Patients were classified as Albustix-negative and
     Albustix-positive patients. The highest urinary alpha-2
     -macroglobulin excretion rates (alpha-2-MER) was found in
     Albustix-positive patients followed by Albustix-negative patients and the
     healthy controls. In view of the fact that the stroke radius of
     alpha-2-macroglobulin (88 ANG ) is larger than
     that of the restrictive pore (56 ANG ), the present finding suggests that
     leakage of alpha-2-macroglobulin in
     urine may be induced by defect of non-discriminatory pores in the
     glomerular basement membrane proposed by Deen and colleagues.
CC
     Radiation biology - Radiation and isotope techniques
     Clinical biochemistry - General methods and applications
     Biochemistry methods - Proteins, peptides and amino acids 10054
     Biochemistry studies - Proteins, peptides and amino acids
     Biochemistry studies - Carbohydrates
                                            10068
     Biophysics - Methods and techniques
                                           10504
     Metabolism - Carbohydrates
                                  13004
     Metabolism - Proteins, peptides and amino acids
                                                       13012
     Metabolism - Metabolic disorders
                                        13020
     Urinary system - Physiology and biochemistry
     Endocrine - Pancreas
                            17008
ΙT
     Major Concepts
        Biochemistry and Molecular Biophysics; Clinical Chemistry (Allied
        Medical Sciences); Endocrine System (Chemical Coordination and
        Homeostasis); Metabolism; Radiology (Medical Sciences); Urinary System
        (Chemical Coordination and Homeostasis)
IT
     Miscellaneous Descriptors
        ANALYTICAL METHOD; CLINICAL FEATURES; DIABETES MELLITUS;
        PROTEINURIA
ORGN Classifier
        Hominidae
                    86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        human
     Taxa Notes
        Animals, Chordates, Humans, Mammals, Primates, Verte
```

```
ANSWER 2 OF 11 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
ΑN
     1995:509170 BIOSIS
     PREV199598514220
DN
TI
     A radioimmunometric assay for urinary alpha-2-
     macroglobulin.
ΑU
     Ito, Seiki; Usami, Akio; Yamazaki, Masatoshi; Shibata, Akira
CS
     Div. Gerontol., Akita Univ. Hosp., Akita 010, Japan
SO
     Tohoku Journal of Experimental Medicine, (1995) Vol. 176, No. 3,
     pp. 137-147.
     CODEN: TJEMAO. ISSN: 0040-8727.
DT
     Article
LA
     English
     Entered STN: 29 Nov 1995
ED
     Last Updated on STN: 29 Nov 1995
AB
     To measure urinary alpha-2-macroglobulin
     levels, a sensitive radioimmunometric assay was established. The least
     detectable level of this assay was 225 pg/ml. A linear correlation
     between alpha-2-macroglobulin levels and
     serial dilution of urine samples was found. Western blot
     analysis and study on column chromatography revealed that the molecular
     weight of alpha-2-macroglobulin in
     urine was identical to that of serum alpha-2-
     macroglobulin. The findings suggested that urinary substance
     detected by the present assay was truly alpha-2-
     macroglobulin. Timed overnight urine samples from 49
     diabetic patients with retinopathy and 20 healthy controls were measured
     by the present assay. Patients were classified as Albustix-negative and
     Albustix-positive patients. The highest urinary alpha-2
     -macroglobulin excretion rates (alpha-2-MER) was found in
     Albustix-positive patients followed by Albustix-negative patients and the
     healthy controls. In view of the fact that the stroke radius of
     alpha-2-macroglobulin (88 ANG ) is larger than
     that of the restrictive pore (56 ANG ), the present finding suggests that
     leakage of alpha-2-macroglobulin in
     urine may be induced by defect of non-discriminatory pores in the
     glomerular basement membrane proposed by Deen and colleagues.
CC
     Radiation biology - Radiation and isotope techniques
     Clinical biochemistry - General methods and applications
                                                                 10006
     Biochemistry methods - Proteins, peptides and amino acids
                                                                 10054
     Biochemistry studies - Proteins, peptides and amino acids
                                                                  10064
     Biochemistry studies - Carbohydrates
                                            10068
     Biophysics - Methods and techniques
                                           10504
     Metabolism - Carbohydrates
                                  13004
     Metabolism - Proteins, peptides and amino acids
                                                       13012
     Metabolism - Metabolic disorders
                                        13020
     Urinary system - Physiology and biochemistry
     Endocrine - Pancreas
                            17008
IT
     Major Concepts
        Biochemistry and Molecular Biophysics; Clinical Chemistry (Allied
        Medical Sciences); Endocrine System (Chemical Coordination and
        Homeostasis); Metabolism; Radiology (Medical Sciences); Urinary System
        (Chemical Coordination and Homeostasis)
IT
     Miscellaneous Descriptors
        ANALYTICAL METHOD; CLINICAL FEATURES; DIABETES MELLITUS;
        PROTEINURIA
ORGN Classifier
        Hominidae 86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        human
     Taxa Notes
        Animals, Chordates, Humans, Mammals, Primates, Verte
```

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(FILE 'HOME' ENTERED AT 11:51:58 ON 15 AUG 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 11:52:24 ON 15 AUG 2007
L1 19581 S URINE AND PEPTIDE?
L2 1122 S L1 AND DIGEST?

1122 S L1 AND DIGEST? L2 247 S L2 AND (MASS SPECTR?) L3 L40 S L3 AND TCA? L5. 0 S L3 AND ACETONE? L6 109 DUPLICATE REMOVE L3 (138 DUPLICATES REMOVED) L7 64 S L6 AND PD<2004 21001 S (ALPHA 2 MACROGLOBULIN) L8 0 S L7 AND L8 L9 L10 3 S L7 AND DIABETE? 363 S L8 AND DIABETE? L11·L12 229 DUPLICATE REMOVE L11 (134 DUPLICATES REMOVED) L13 . 193 S L12 AND PD<2004 0 S L13 AND (MASS SPECTRO?) L14 ·

L15 8 S L13 AND URINE?

L16 9 S L13 AND PEPTIDE? L17 9 S L16 NOT L15

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=>

(FILE 'HOME' ENTERED AT 11:51:58 ON 15 AUG 2007)

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FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 11:52:24 ON 15
     AUG 2007
L1
         19581 S URINE AND PEPTIDE?
          1122 S L1 AND DIGEST?
· L2
           247 S L2 AND (MASS SPECTR?)
L3
              0 S L3 AND TCA?
L4 .
              0 S L3 AND ACETONE?
L5
_{
m L6}
          . 109 DUPLICATE REMOVE L3 (138 DUPLICATES REMOVED)
L7
             64 S L6 AND PD<2004
rs
          21001 S (ALPHA 2 MACROGLOBULIN)
              0 S L7 AND L8
L9
              3 S L7 AND DIABETE?
L10
         . 363 S L8 AND DIABETE?
L11
            229 DUPLICATE REMOVE L11 (134 DUPLICATES REMOVED)
L12
L13
            193 S L12 AND PD<2004
             0 S L13 AND (MASS SPECTRO?)
L14
             8 S L13 AND URINE?
L15
L16
             9 S L13 AND PEPTIDE?
             9 S L16 NOT L15
L17
```

```
ΑN
     2002:778627 CAPLUS
     137:259345
DN
ED
     Entered STN: 11 Oct 2002
TТ
     Method for the quantitative determination of proteinase inhibitors in the
     body fluids of human or animal using porcine pancreatic elastase and
     diagnostic applications
ΙN
     Bristow, Cindy L.
PΑ
     USA
     U.S. Pat. Appl. Publ., 11 pp., Cont.-in-part of U.S. Ser. No. 452,699.
SO
     CODEN: USXXCO
DT
     Patent
LA
     English
IC
     ICM C120001-37
     ICS G06F019-00; G01N033-48; G01N033-50
INCL 435023000; X70-2 1.9
     7-3 (Enzymes)
     Section cross-reference(s): 14
FAN.CNT 2
     PATENT NO.
                         KIND
                                 DATE
                                             APPLICATION NO.
                                                                     DATE
                         ____
     US 2002146756
                          A1
                                 20021010
ΡI
                                             US 2002-105719
                                                                     20020325 <--
     US 6887678
                          В2
                                 20050503
                        P
PRAI US 1998-110580P
                                 19981202
     US 1999-452699
                         A2
                                 19991202
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 US 2002146756
                 ICM
                        C12Q001-37
                 ICS
                        G06F019-00; G01N033-48; G01N033-50
                 INCL.
                        435023000; X70-2 1.9
                 IPCI
                        C12Q0001-37 [ICM,7]; G06F0019-00 [ICS,7]; G01N0033-48
                         [ICS, 7]; G01N0033-50 [ICS, 7]
                 IPCR
                        C12Q0001-37 [I,C*]; C12Q0001-37 [I,A]
                 NCL
                        435/023.000; 702/019.000
                 ECLA
                        C12Q001/37
AB
     A method is provided for the quant. detns. of active and inactive concns.
     of proteinase inhibitors, such as \alpha 1-antitrypsin (\alpha 1PI) and .
     alpha.2-macroglobulin (\alpha 2M), in the body
     fluids of humans and animals. Porcine pancreatic elastase were used in
     the assays for \alpha 1PI and \alpha 2M. Diagnostic applications of the
     method are presented.
     proteinase inhibitor detn elastase body fluid diagnosis; elastase alphal
ST
     antitrypsin alphal alpha2 macroglobulin detn diagnosis
ΙT
     Infection
        (bacterial; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     Diagnosis
        (cancer; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     AIDS (disease)
        (infection; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     Autoimmune disease
        (insulin-dependent diabetes mellitus; method for quant. determination
        of proteinase inhibitors in body fluids of human or animal using
        porcine pancreatic elastase and diagnostic applications)
IT
     Diabetes mellitus
        (insulin-dependent; method for quant. determination of proteinase
inhibitors in
        body fluids of human or animal using porcine pancreatic elastase and
```

ANSWER 3 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN

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ANSWER 3 OF 8 CAPLUS COPYRIGHT 2007 ACS on STN
     2002:778627 CAPLUS
ΑN
     137:259345
DN
     Entered STN: 11 Oct 2002
ED
TΤ
     Method for the quantitative determination of proteinase inhibitors in the
     body fluids of human or animal using porcine pancreatic elastase and
     diagnostic applications
ΙN
     Bristow, Cindy L.
PΑ
     USA
SO
     U.S. Pat. Appl. Publ., 11 pp., Cont.-in-part of U.S. Ser. No. 452,699.
     CODEN: USXXCO
DT
     Patent
LA
     English
IC
     ICM C12Q001-37
     ICS G06F019-00; G01N033-48; G01N033-50
INCL 435023000; X70-2 1.9
     7-3 (Enzymes)
     Section cross-reference(s): 14
FAN.CNT 2
     PATENT NO.
                                      APPLICATION NO.
                        KIND DATE
                                                                  DATE
                       ____
                               -----
                                           -----
                                                                  -----
                        A1
     US 2002146756
                                20021010
                                          US 2002-105719
                                                                  20020325 <--
                        B2
     US 6887678
                                20050503
PRAI US 1998-110580P P US 1999-452699 A2
                               19981202
                                19991202
CLASS
 PATENT NO.
                 CLASS PATENT FAMILY CLASSIFICATION CODES
 ______
                ____
                 ICM
                        C12Q001-37
 US 2002146756
                 ICS
                        G06F019-00; G01N033-48; G01N033-50
                 INCL
                        435023000; X70-2 1.9
                        C12Q0001-37 [ICM,7]; G06F0019-00 [ICS,7]; G01N0033-48
                 IPCI
                        [ICS, 7]; G01N0033-50 [ICS, 7]
                 IPCR
                        C12Q0001-37 [I,C*]; C12Q0001-37 [I,A]
                 NCL
                        435/023.000; 702/019.000
                 ECLA
                        C12Q001/37
     A method is provided for the quant. detns. of active and inactive concns.
AB
     of proteinase inhibitors, such as \alpha 1-antitrypsin (\alpha 1PI) and .
     alpha.2-macroglobulin (\alpha 2M), in the body
     fluids of humans and animals. Porcine pancreatic elastase were used in
     the assays for \alpha 1PI and \alpha 2M. Diagnostic applications of the
     method are presented.
     proteinase inhibitor detn elastase body fluid diagnosis; elastase alphal
ST
     antitrypsin alphal alpha2 macroglobulin detn diagnosis
IT
     Infection
        (bacterial; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
TТ
     Diagnosis
        (cancer; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
·IT
     AIDS (disease)
        (infection; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     Autoimmune disease
        (insulin-dependent diabetes mellitus; method for quant. determination
        of proteinase inhibitors in body fluids of human or animal using
        porcine pancreatic elastase and diagnostic applications)
IT
     Diabetes mellitus
        (insulin-dependent; method for quant. determination of proteinase
inhibitors in
        body fluids of human or animal using porcine pancreatic elastase and
```

```
diagnostic applications)
IT
     Aging, animal
     Animals
     Arthritis
     Ascitic fluid
     Asthma
     Atherosclerosis
     Blood analysis
     Body fluid
     Human
     Lymph node, disease
     Neoplasm
     Periodontium, disease
     Regression analysis
     Saliva
     Tear (ocular fluid)
       Urine analysis
         (method for quant. determination of proteinase inhibitors in body fluids of
        human or animal using porcine pancreatic elastase and diagnostic
        applications)
IT
     Diagnosis
        (mol.; method for quant. determination of proteinase inhibitors in body
fluids
        of human or animal using porcine pancreatic elastase and diagnostic
        applications)
IT
     Nose
        (nasal specimens; method for quant. determination of proteinase inhibitors
in
        body fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
TΤ
     Enzyme kinetics
        (of inhibition; method for quant. determination of proteinase inhibitors in
        body fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
·IT
     Infection
        (parasitic; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
ΙŢ
        (plasma; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     Lupus erythematosus
        (systemic; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     Vagina
        (vaginal specimens; method for quant. determination of proteinase
inhibitors in
        body fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
IT
     Infection
        (viral; method for quant. determination of proteinase inhibitors in body
fluids
        of human or animal using porcine pancreatic elastase and diagnostic
        applications)
ΙT
     Macroglobulins
     RL: ANT (Analyte); DGN (Diagnostic use); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        (\alpha 2-; method for quant. determination of proteinase inhibitors in body
        fluids of human or animal using porcine pancreatic elastase and
        diagnostic applications)
     9041-92-3, \alpha1-Antitrypsin
                                  37205-61-1, Proteinase inhibitor
     RL: ANT (Analyte); DGN (Diagnostic use); ANST (Analytical study); BIOL
```

```
diagnostic applications)
 ΙT
      Aging, animal
      Animals
      Arthritis
      Ascitic fluid
      Asthma
      Atherosclerosis
      Blood analysis
      Body fluid
      Human
      Lymph node, disease
      Neoplasm
      Periodontium, disease
      Regression analysis
      Saliva
      Tear (ocular fluid)
        Urine analysis
         (method for quant. determination of proteinase inhibitors in body fluids of
         human or animal using porcine pancreatic elastase and diagnostic
         applications)
 ΙT
      Diagnosis
         (mol.; method for quant. determination of proteinase inhibitors in body
 fluids
         of human or animal using porcine pancreatic elastase and diagnostic
         applications)
·IT
      Nose
         (nasal specimens; method for quant. determination of proteinase inhibitors
in
         body fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
·IT
      Enzyme kinetics
         (of inhibition; method for quant. determination of proteinase inhibitors in
         body fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
IT
      Infection
         (parasitic; method for quant. determination of proteinase inhibitors in body
         fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
ΤТ
         (plasma; method for quant. determination of proteinase inhibitors in body
         fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
ΙT
      Lupus erythematosus
         (systemic; method for quant. determination of proteinase inhibitors in body
         fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
ΙT
      Vagina
         (vaginal specimens; method for quant. determination of proteinase
inhibitors in
        body fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
IT
      Infection
         (viral; method for quant. determination of proteinase inhibitors in body
fluids
         of human or animal using porcine pancreatic elastase and diagnostic
         applications)
\cdot IT
      Macroglobulins
      RL: ANT (Analyte); DGN (Diagnostic use); ANST (Analytical study); BIOL
      (Biological study); USES (Uses)
         (\alpha 2-; method for quant. determination of proteinase inhibitors in body
         fluids of human or animal using porcine pancreatic elastase and
         diagnostic applications)
ΙT
      9041-92-3, \alpha1-Antitrypsin
                                 37205-61-1, Proteinase inhibitor
      RL: ANT (Analyte); DGN (Diagnostic use); ANST (Analytical study); BIOL
```

(Biological study); USES (Uses)

(method for quant. determination of proteinase inhibitors in body fluids of human or animal using porcine pancreatic elastase and diagnostic applications)

IT 9004-06-2, Elastase

RL: ARG (Analytical reagent use); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(method for quant. determination of proteinase inhibitors in body fluids of human or animal using porcine pancreatic elastase and diagnostic applications)

RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Anon; SU 1573430 A 1990 CAPLUS
- (2) Anon; DE 3938971 Al 1991 CAPLUS
- (3) Anon; RU 2039983 C1 1995 CAPLUS
- (4) Anon; EP 0288841 A2 1998 CAPLUS
- (5) Bristow, C; Clinical and Diagnostic Lab Immunology. 2001, V8(5), P937 MEDLINE
- (6) Coan; US 4697003 A 1987 CAPLUS
- (7) Lloyd; US 5073487 A 1991 CAPLUS
- (8) Ralston; US 6093804 A 2000 CAPLUS
- (9) Simon; US 5773430 A 1998 CAPLUS
- (10) Travis; US 4493891 A 1985 CAPLUS

(Biological study); USES (Uses)

(method for quant. determination of proteinase inhibitors in body fluids of human or animal using porcine pancreatic elastase and diagnostic applications)

IT 9004-06-2, Elastase

RL: ARG (Analytical reagent use); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(method for quant. determination of proteinase inhibitors in body fluids of human or animal using porcine pancreatic elastase and diagnostic applications)

RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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- (6) Coan; US 4697003 A 1987 CAPLUS
- (7) Lloyd; US 5073487 A 1991 CAPLUS
- (8) Ralston; US 6093804 A 2000 CAPLUS
- (9) Simon; US 5773430 A 1998 CAPLUS
- ·(10) Travis; US 4493891 A 1985 CAPLUS.

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ANSWER 8 OF 9
                  MEDLINE on STN
     2002133559
                    MEDLINE
     PubMed ID: 11868856
     A study of plasma alpha-2-macroglobulin
     levels in type 2 diabetic subjects with microalbuminuria.
ΑU
     Ahmad J; Singh M; Saleemuddin M
CS
     Department of Medicine, JN Medical College, Aligarh.
SO
     The Journal of the Association of Physicians of India, (2001 Nov)
     Vol. 49, pp. 1062-5.
     Journal code: 7505585. ISSN: 0004-5772.
CY
DT
     (CLINICAL TRIAL)
     (COMPARATIVE STUDY)
     Journal; Article; (JOURNAL ARTICLE)
     (RANDOMIZED CONTROLLED TRIAL)
     (RESEARCH SUPPORT, NON-U.S. GOV'T)
LA
     English
FS
     Priority Journals
EM
     200203
     Entered STN: 1 Mar 2002
     Last Updated on STN: 15 Mar 2002
     Entered Medline: 14 Mar 2002
AB
     BACKGROUND: Alpha-2 macroglobulin
     (Alpha-2-M) is a major plasma protease inhibitor that also regulates the
     activity of a variety of bioactive peptides including
     interleukins and exerts a range of immunomodulatory effects.
     We conducted the present study with the objective to study the alpha-2-M
     levels in type 2 diabetic subjects with microalbuminuria in an attempt to
     establish alpha-2-M as a predictor of microvascular complications in
     diabetes. MATERIAL AND METHODS: Plasma Alpha-2-M levels were
     assayed in 100 (53 males and 47 females) randomly selected type 2 diabetic
     subjects with microalbuminuria. Diabetes was diagnosed
     according to the expert committee report of 1998. Patients with any acute
     metabolic complication like hypoglycemia, ketoacidosis, cerebrovascular
     accident or any acute infection were not included in the study group.
     RESULTS: Majority of patients belonged to 40-60 years age group. In our
     study alpha-2-M levels indicated a clear increase in diabetic subjects
     with the increasing age of subjects confirmed by multiple logistical
     analysis. Alpha-2-M levels were not found to be significantly different
     between males and females (55.6 \pm/- 11.3 vs. 53.7 \pm/- 10.5). Duration of
     diabetes was found to be an important confounding variable showing
     a direct positive correlation with alpha-2-M levels and also a significant
     correlation was found between alpha-2-M levels with different levels of
     microalbuminuria on multiple logistical analysis. No significant relation
     of alpha-2-M levels with either fasting blood sugar or HbA1 was observed.
     CONCLUSION: The increase in plasma alpha-2
     macroglobulin levels in diabetes may be a correlative
     measure to encounter the potential proteolytic challenge associated with
     diabetic microangiopathy, even very early in the course of the disease.
     Alph-2 macroglobulin may yet be one of the most specific markers of
     microvascular complications in diabetes than any other serum
     protein.
     Check Tags: Female; Male
      Aged
     *Albuminuria: DI, diagnosis
       *Diabetes Mellitus, Type 2: BL, blood
        Diabetes Mellitus, Type 2: DI, diagnosis
      Diabetic Angiopathies: DI, diagnosis
      Humans
      Logistic Models
      Middle Aged
      Predictive Value of Tests
      Probability
      Prognosis
```

```
MEDLINE on STN
     2002133559
                    MEDLINE
     PubMed ID: 11868856
     A study of plasma alpha-2-macroglobulin
     levels in type 2 diabetic subjects with microalbuminuria.
ΑU
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     Department of Medicine, JN Medical College, Aligarh.
SO
     The Journal of the Association of Physicians of India, (2001 Nov)
     Vol. 49, pp. 1062-5.
     Journal code: 7505585. ISSN: 0004-5772.
.CY
DT
     (CLINICAL TRIAL)
     (COMPARATIVE STUDY)
     Journal; Article; (JOURNAL ARTICLE)
     (RANDOMIZED CONTROLLED TRIAL)
     (RESEARCH SUPPORT, NON-U.S. GOV'T)
LA
     English
FS
     Priority Journals
EM
     200203
ĒD
     Entered STN: 1 Mar 2002
     Last Updated on STN: 15 Mar 2002
     Entered Medline: 14 Mar 2002
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     activity of a variety of bioactive peptides including
     interleukins and exerts a range of immunomodulatory effects.
                                                                    OBJECTIVE:
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     diabetic microangiopathy, even very early in the course of the disease.
     Alph-2 macroglobulin may yet be one of the most specific markers of
     microvascular complications in diabetes than any other serum
     protein.
CT
     Check Tags: Female; Male
      Aged
     *Albuminuria: DI, diagnosis
       *Diabetes Mellitus, Type 2: BL, blood
        Diabetes Mellitus, Type 2: DI, diagnosis
      Diabetic Angiopathies: DI, diagnosis
      Humans
      Logistic Models
      Middle Aged
      Predictive Value of Tests
      Probability
      Prognosis
```

ANSWER 8 OF 9

Prospective Studies
Sensitivity and Specificity
Severity of Illness Index
*alpha-Macroglobulins: AN, analysis
CN 0 (alpha-Macroglobulins)

Prospective Studies
Sensitivity and Specificity
Severity of Illness Index
*alpha-Macroglobulins: AN, analysis
CN 0 (alpha-Macroglobulins)

10/570,836 Search Lycook 8/15/07

d his

(FILE 'HOME' ENTERED AT 14:51:32 ON 15 AUG 2007)

| FILE 'BIOSIS, | CAPLUS, | EMBASE, | MEDLINE, | JAPIO' | ENTERED | ΑT | 14:51:52 | ON | 15 |
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| AUG 2007 | | | | | | | | | |

| | AUG | 2007 | |
|----|-----|-------|----------------------------------------------|
| L1 | | 175 | S (ALPHA 2 MACROGLOBULIN) AND (MASS SPECTR?) |
| L2 | | 100 | DUPLICATE REMOVE L1 (75 DUPLICATES REMOVED) |
| L3 | | 42 | S L2 AND PD<2004 |
| L4 | | 0 | S L3 AND DIAB? |
| L5 | | 8 | S L3 AND DIGEST? |
| L6 | | 21003 | S (ALPHA 2 MACROGLOBULIN) |
| L7 | | 13 | S L6 AND FINGERPRINT? |
| L8 | | 6 | DUPLICATE REMOVE L7 (7 DUPLICATES REMOVED) |
| L9 | | 4 | S L8 AND PD<2004 |

(FILE 'HOME' ENTERED AT 14:51:32 ON 15 AUG 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 14:51:52 ON 15 AUG 2007

- L1 175 S (ALPHA 2 MACROGLOBULIN) AND (MASS SPECTR?)
- L2 100 DUPLICATE REMOVE L1 (75 DUPLICATES REMOVED)
- L3 42 S L2 AND PD<2004
- L4 0 S L3 AND DIAB?
 - 8 S L3 AND DIGEST?
- L6 21003 S (ALPHA 2 MACROGLOBULIN)
- L7 13 S L6 AND FINGERPRINT?
- L8 6 DUPLICATE REMOVE L7 (7 DUPLICATES REMOVED)
- L9 4 S L8 AND PD<2004

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L5

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ANSWER 1 OF 4 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
     2004:78571 BIOSIS
     PREV200400081442
DN
ΤI
     Elevated levels of serum alpha2 macroglobulin in wild black bears during
ΑU
     Sheikh, Ashfaq M.; Chauhan, Ved; Tsiouris, John A. [Reprint Author];
     Mehta, Pankaj D.; Burguess, Kelcey; Fenko, Michael D.; Spivack, Warren;
     Vaughan, Michael; Malik, Mazhar
     NYS Institute for Basic Research in Developmental Disabilities, 1050
CS
     Forest Hill Road, Staten Island, NY, 10314, USA
     john.Tsiouris@omr.state.ny.us
SO
     Biochimie (Paris), (October 2003) Vol. 85, No. 10, pp.
     1027-1032. print.
     CODEN: BICMBE. ISSN: 0300-9084.
DT
     Article
     English
LA
ED
     Entered STN: 4 Feb 2004
     Last Updated on STN: 4 Feb 2004
     Bear serum alpha2 macroglobulin (alpha2M) was purified by sodium dodecyl
AΒ
     sulfate polyacrylamide gel electrophoresis (SDS-PAGE) and partially
     characterized by tryptic digestion of alpha2M and analysis of the peptides
     by peptide mass fingerprinting. The molecular weight of bear
     serum alpha2M was 181 kDa, same as for human serum alpha2M, on SDS-PAGE.
     However, the MALDI mass spectrum of the tryptic digested bear serum
     alpha2M showed that it is different from human alpha2M or other data bank
     proteins. Liquid chromatography (LC)/mass spectrometry (MS)/MS of the
     proteolytic products of bear serum alpha2M showed eight peptides that had
     similarities to human alpha2M suggesting that the protein of interest was
     indeed alpha2M of bear. The polyclonal antibody against bear serum
     alpha2M recognized only one protein from the western blot of bear serum
     proteins. It also recognized human alpha2M. The levels of serum alpha2M
     were significantly increased during hibernating state as compared to
     active state of bears indicating its protective role from the consequences
     of the metabolic depression during hibernation.
     Biochemistry studies - General
                                      10060
     Enzymes - General and comparative studies: coenzymes
                                                             10802
     Blood - Blood and lymph studies
                                       15002
     Blood - Blood cell studies
ΙT
     Major Concepts
        Biochemistry and Molecular Biophysics
IT
     Parts, Structures, & Systems of Organisms
        serum: blood and lymphatics
IT
     Chemicals & Biochemicals
          alpha-2 macroglobulin: characterization,
        purification; trypsin [EC 3.4.21.4]
ΙT
     Miscellaneous Descriptors
        hibernation; metabolic depression
ORGN Classifier
        Hominidae
                    86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        human (common)
     Taxa Notes
        Animals, Chordates, Humans, Mammals, Primates, Vertebrates
ORGN Classifier
        Ursidae
                  85790
     Super Taxa
        Carnivora; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        black bear (common): wild
     Taxa Notes
        Animals, Carnivores, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman
        Mammals, Vertebrates
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ANSWER 1 OF 4 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
AN
     2004:78571 BIOSIS
DN
     PREV200400081442
ΤI
     Elevated levels of serum alpha2 macroglobulin in wild black bears during
ΑU
     Sheikh, Ashfaq M.; Chauhan, Ved; Tsiouris, John A. [Reprint Author];
     Mehta, Pankaj D.; Burguess, Kelcey; Fenko, Michael D.; Spivack, Warren;
     Vaughan, Michael; Malik, Mazhar
CS
     NYS Institute for Basic Research in Developmental Disabilities, 1050
     Forest Hill Road, Staten Island, NY, 10314, USA
     john.Tsiouris@omr.state.ny.us
SO
     Biochimie (Paris), (October 2003) Vol. 85, No. 10, pp.
     1027-1032. print.
     CODEN: BICMBE. ISSN: 0300-9084.
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     Article
LA
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     Last Updated on STN: 4 Feb 2004
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     However, the MALDI mass spectrum of the tryptic digested bear serum
     alpha2M showed that it is different from human alpha2M or other data bank
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     alpha2M recognized only one protein from the western blot of bear serum
     proteins. It also recognized human alpha2M. The levels of serum alpha2M
     were significantly increased during hibernating state as compared to
     active state of bears indicating its protective role from the consequences
     of the metabolic depression during hibernation.
     Biochemistry studies - General
                                      10060
     Enzymes - General and comparative studies: coenzymes
                                                             10802
     Blood - Blood and lymph studies
                                       15002
     Blood - Blood cell studies 15004
ΙT
     Major Concepts
        Biochemistry and Molecular Biophysics
IT
     Parts, Structures, & Systems of Organisms
        serum: blood and lymphatics
IT
     Chemicals & Biochemicals
          alpha-2 macroglobulin: characterization,
        purification; trypsin [EC 3.4.21.4]
IT
     Miscellaneous Descriptors
        hibernation; metabolic depression
ORGN Classifier
        Hominidae
                    86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        human (common)
     Taxa Notes
        Animals, Chordates, Humans, Mammals, Primates, Vertebrates
ORGN Classifier
        Ursidae
                  85790
     Super Taxa
        Carnivora; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        black bear (common): wild
     Taxa Notes
        Animals, Carnivores, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman
        Mammals, Vertebrates
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RN 9002-07-7 (trypsin) 9002-07-7 (EC 3.4.21.4) RN 9002-07-7 (trypsin) 9002-07-7 (EC 3.4.21.4) d his

(FILE 'HOME' ENTERED AT 18:34:36 ON 15 AUG 2007)

| | FILE 'BIOS' | IS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 18:34:56 ON 15 |
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| L1 | 588464 | S (MASS SPECTROMET?) |
| L2 | 60313 | S L1 AND PEPTIDE? |
| L3 | 1059 | S L2 AND URINE? |
| L4 | 1 | S L3 AND TCA? |
| L5 | 226 | S L3 AND ELECTROPHORESIS? |
| L6 | 149 | DUPLICATE REMOVE L5 (77 DUPLICATES REMOVED) |
| L7 | 54 | S L6 AND PD<2004 |
| L8 | 5 | S L7 AND DIABETE? |
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(FILE 'HOME' ENTERED AT 18:34:36 ON 15 AUG 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, JAPIO' ENTERED AT 18:34:56 ON 15 AUG 2007

L1 588464 S (MASS SPECTROMET?)

L2 60313 S L1 AND PEPTIDE?

L3 1059 S L2 AND URINE?

L4 1 S L3 AND TCA?

L5 226 S L3 AND ELECTROPHORESIS?

L6 149 DUPLICATE REMOVE L5 (77 DUPLICATES REMOVED)

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ANSWER 17 OF 17 CAPLUS COPYRIGHT 2007 ACS on STN
     2002:869179 CAPLUS
AN
     137:366001
DN
ED
     Entered STN: 15 Nov 2002
     Process for preparation and analysis of protein samples
TI
     Parker, Kenneth C.; Nadler, Timothy K.; Vella, George J.; Huang, Yulin;
IN
     Aebersold, Rudolf H.; Smolka, Marcus B.
PA
     Perseptive Biosystems, Inc., USA; Institute for Systems Biology
SO
     PCT Int. Appl., 39 pp.
     CODEN: PIXXD2
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IC
CC
     9-16 (Biochemical Methods)
     Section cross-reference(s): 6
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                                   DATE
                           KIND
                                                APPLICATION NO.
                                                                          DATE
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                       A2
     WO 2002090929
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                          G01N033/68A
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                  ECLA
                         G01N033/68A
AΒ
     The invention concerns methods using gel electrophoresis and
     mass spectrometry for the rapid, quant. anal. of
     proteins or protein function in mixts. of proteins derived from two or
     more samples in one unit operation. In one embodiment the method includes
     (a) preparing an extract of proteins from each of at least two different
     samples; (b) providing a set of substantially chemical identical and
     differentially isotopically labeled protein reagents, one for each sample;
     (c) reacting each protein sample of step (a) with a different reagent from
     the set of step (b) to provide isotopically labeled proteins; (d) mixing
     each of said isotopically labeled proteins to form a single mixture of
     different isotopically labeled proteins; (e) electrophoresing the mixture of
     step (d) by an electrophoresing method capable of separating proteins within
     said mixture; and (f) detecting the difference in the expression levels of
     the proteins in the two samples by spectrometry based on individual
     peptides derived from chemical or enzymic digestion.
     anal. method can be used for qual. and particularly for quant. anal. of
     global protein expression profiles in cells and tissues, i.e. the quant.
     anal. of proteomes.
ST
     protein sample prep gel electrophoresis mass
     spectrometry digestion label
IT
     Reagents
     RL: NUU (Other use, unclassified); USES (Uses)
         (ICAT; process for preparation and anal. of protein samples)
IT
     Enzymes, uses
     RL: NUU (Other use, unclassified); USES (Uses)
        (for digestion; process for preparation and anal. of protein
        samples)
ΙT
     Organelle
         (membrane-containing; process for preparation and anal. of protein samples)
ΙT
     Animal cell
     Animal tissue
     Ascites
     Blood serum .
```

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G01N0033-68 [ICS, 7]
                  IPCR
                         G01N0027-62 [I,C*]; G01N0027-447 [I,C*]; G01N0033-48
                         [I,C*]; G01N0033-483 [I,C*]; G01N0033-50 [I,C*];
                         G01N0033-60 [I,C*]; G01N0033-68 [I,C*]; G01N0027-62 [I,A]; G01N0027-447 [I,A]; G01N0033-48 [I,A];
                         G01N0033-483 [I,A]; G01N0033-50 [I,A]; G01N0033-60
                         [I,A]; G01N0033-68 [I,A]
 EP 1392848
                  IPCI
                         G01N0033-68 [I,C]; G01N0033-60 [I,C]; G01N0033-68
                         [I,A]; G01N0033-60 [I,A]
                         G01N0027-62 [I,C*]; G01N0027-62 [I,A]; G01N0033-68 [I,C]; G01N0033-68 [I,A]; G01N0027-447 [I,C*];
                  IPCR ·
                         G01N0027-447 [I,A]; G01N0033-48 [I,C*]; G01N0033-48 [I,A]; G01N0033-483 [I,C*]; G01N0033-483 [I,A];
                         G01N0033-50 [I,C*]; G01N0033-50 [I,A]; G01N0033-60
                         [I,C]; G01N0033-60 [I,A]
                  ECLA
                         G01N033/68A
 JP 2004533610
                  IPCI
                         G01N0033-483 [ICM,7]; G01N0027-62 [ICS,7]; G01N0033-48
                          [ICS,7]; G01N0033-50 [ICS,7]; G01N0027-447 [ICS,7]
                  IPCR
                         G01N0033-60 [I,A]; G01N0033-60 [I,C*]; G01N0033-68
                         [I,A]; G01N0033-68 [I,C*]
                  FTERM
                         2G045/AA34; 2G045/BA13; 2G045/BB03; 2G045/BB14;
                         2G045/BB51; 2G045/CB01; 2G045/DA36; 2G045/FB05;
                         2G045/FB06; 2G045/FB08; 2G045/JA01
 AT 350666
                  IPCI
                         G01N0033-68 [ICS,7]; G01N0033-60 [ICS,7]
                         G01N0027-62 [I,C*]; G01N0027-447 [I,C*]; G01N0033-48
                  IPCR
                         [I,C*]; G01N0033-483 [I,C*]; G01N0033-50 [I,C*];
                         G01N0033-60 [I,C*]; G01N0033-68 [I,C*]; G01N0027-62
                         [I,A]; G01N0027-447 [I,A]; G01N0033-48 [I,A];
                         G01N0033-483 [I,A]; G01N0033-50 [I,A]; G01N0033-60
                         [I,A]; G01N0033-68 [I,A]
                         G01N033/68A
                  ECLA
AB
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     mass spectrometry for the rapid, quant. anal. of
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     spectrometry digestion label
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     RL: NUU (Other use, unclassified); USES (Uses)
         (for digestion; process for preparation and anal. of protein
        samples)
IT
     Organelle
         (membrane-containing; process for preparation and anal. of protein samples)
ΙT
     Animal cell
     Animal tissue
     Ascites
     Blood serum
```

```
Cell nucleus
    Cerebrospinal fluid
       Digestion, biological
     Gel electrophoresis
    Labels
      Mass spectrometry
     Post-translational processing
     Protein motifs
     Sample preparation
     Separation
     Staining, coloring
        (process for preparation and anal. of protein samples)
ΙT
     Proteins
     RL: ANT (Analyte); PUR (Purification or recovery); ANST (Analytical
     study); PREP (Preparation)
        (process for preparation and anal. of protein samples)
IT
     Isotopes
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (process for preparation and anal. of protein samples)
ΙT
     475134-24-8
                   475134-25-9
                                 475134-26-0
     RL: PRP (Properties)
        (unclaimed sequence; process for preparation and anal. of protein samples)
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=>

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Cell nucleus
    Cerebrospinal fluid
       Digestion, biological
    Gel electrophoresis
    Labels
       Mass spectrometry
     Post-translational processing
     Protein motifs
     Sample preparation
     Separation
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        (process for preparation and anal. of protein samples)
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     Proteins
     RL: ANT (Analyte); PUR (Purification or recovery); ANST (Analytical
     study); PREP (Preparation)
        (process for preparation and anal. of protein samples)
ΙT
     Isotopes
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (process for preparation and anal. of protein samples)
ΙT
     475134-24-8
                   475134-25-9
                                 475134-26-0
     RL: PRP (Properties)
        (unclaimed sequence; process for preparation and anal. of protein samples)
```

ANSWER 2 OF 17 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

AN 2001:415170 BIOSIS

DN PREV200100415170

TI Towards defining the urinary proteome using liquid chromatography-tandem mass spectrometry: I. Profiling an unfractionated tryptic digest.

AU Spahr, Chris S.; Davis, Michael T.; McGinley, Michael D.; Robinson, John H.; Bures, Edward J.; Beierle, Jill; Mort, Jessica; Courchesne, Paul L.; Chen, Kui; Wahl, Robert C.; Yu, Wen; Luethy, Roland; Patterson, Scott D. [Reprint author]

CS Celera Genomics, 45 West Gude Drive, Rockville, MD, 20850, USA scott.patterson@celera.com

SO Proteomics, (January, 2001) Vol. 1, No. 1, pp. 93-107. print. ISSN: 1615-9853.

DT Article

LA English

ED Entered STN: 29 Aug 2001 Last Updated on STN: 22 Feb 2002

AB The proteome of normal male urine from a commercial pooled source has been examined using direct liquid chromatography-tandem mass spectrometry (LC-MS/MS). The entire urinary protein mixture was denatured, reduced and enzymatically digested prior to LC-MS/MS analysis using a hybrid-quadrupole time-of-flight mass spectrometer (Q-TOF) to perform data-dependent ion selection and fragmentation. To fragment as many peptides as possible, the mixture was analyzed four separate times, with the mass spectrometer selecting ions for fragmentation from a subset of the entire mass range for each run. This approach requires only an autosampler on the HPLC for automation (i. e, unattended operation). Across these four analyses, 1.450 peptide MS/MS spectra were matched to 751 sequences to identify 124 gene products (proteins and translations of expressed sequence tags). Interestingly, the experimental time for these analyses was less than that required to run a single two-dimensional gel.

CC Biochemistry studies - General 10060

IT Major Concepts

ΙT

Biochemistry and Molecular Biophysics; Methods and Techniques Chemicals & Biochemicals

human urinary protein: Sigma, lyophilized; unfractionated tryptic digest: profiling; urinary proteome: definition

IT Methods & Equipment

HP 1100 HPLC system [HP 1100 high performance liquid chromatography system]: Hewlett-Packard, equipment; Investigator 2-D electrophoresis system: ESA, equipment; Micromass hybrid quadrupole-time of flight mass spectrometer: Micromass, equipment; hybrid-quadrupole time-of-flight mass spectrometer: equipment; liquid chromatography-tandem mass spectrometry: analytical method, comparison, liquid chromatography, spectroscopy: CB; two-dimensional gel electrophoresis: comparison, polyacrylamide gel electrophoresis, separation method

IT Miscellaneous Descriptors

peptide fragmentation; proteomics

ANSWER 2 OF 17 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

AN 2001:415170 BIOSIS

DN PREV200100415170

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- AU Spahr, Chris S.; Davis, Michael T.; McGinley, Michael D.; Robinson, John H.; Bures, Edward J.; Beierle, Jill; Mort, Jessica; Courchesne, Paul L.; Chen, Kui; Wahl, Robert C.; Yu, Wen; Luethy, Roland; Patterson, Scott D. [Reprint author]
- CS Celera Genomics, 45 West Gude Drive, Rockville, MD, 20850, USA scott.patterson@celera.com
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- CC Biochemistry studies General 10060
- IT Major Concepts

ΙT

Biochemistry and Molecular Biophysics; Methods and Techniques

Chemicals & Biochemicals

human urinary protein: Sigma, lyophilized; unfractionated tryptic digest: profiling; urinary proteome: definition

IT Methods & Equipment

HP 1100 HPLC system [HP 1100 high performance liquid chromatography system]: Hewlett-Packard, equipment; Investigator 2-D electrophoresis system: ESA, equipment; Micromass hybrid quadrupole-time of flight mass spectrometer:
Micromass, equipment; hybrid-quadrupole time-of-flight mass spectrometer: equipment; liquid chromatography-tandem mass spectrometry: analytical method, comparison, liquid chromatography, spectroscopy: CB; two-dimensional gel electrophoresis: comparison, polyacrylamide gel electrophoresis, separation method

IT Miscellaneous Descriptors

peptide fragmentation; proteomics

ANSWER 1 OF 17 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN ΑN 2002:566278 BIOSIS PREV200200566278 DN TТ Proteomics of renal disorders: Urinary proteome analysis by two-dimensional gel electrophoresis and MALDI-TOF mass spectrometry. Kumar, Yadunanda; Uppuluri, Nageshwar Rao Venkata; Babu, Kishore; Phadke, ΑIJ Kishore; Kumar, Prasanna; Ballal, Sudarshan; Tatu, Utpal [Reprint author] CS Department of Biochemistry, Indian Institute of Science, Bangalore, 560 012, India tatu@biochem.iisc.ernet.in Current Science (Bangalore), (25 March, 2002) Vol. 82, No. 6, SO pp. 655-663. print. CODEN: CUSCAM. ISSN: 0011-3891. DT Article LA English Entered STN: 7 Nov 2002 ED Last Updated on STN: 7 Nov 2002 AB The proteomes of urinary samples from patients with different renal conditions were analysed by two-dimensional electrophoresis and MALDI-TOF technology. Samples from three different renal conditions, namely kidney failure, nephrotic syndrome and microalbuminuria, were included in the analysis. Apart from the presence of albumin, the profiles of protein spots found in these urine samples were quite distinct. While kidney failure patients showed predominantly low molecular weight proteins, the nephrotic syndrome patients showed an abundance of relatively high molecular weight proteins clustering in the acidic range of the 2-D gels. Two different protein spots from kidney failure patients, four from nephrotic syndrome patients and three from micro-albuminuria patients were identified by in-gel protease digestions and analysis of resulting peptides by MALDI-TOF. The proteins identified were albumin, alpha-1-antitrypsin, alpha-1-acid glycoprotein 2, Zn-alpha-2-glycoprotein and alpha-1-microglobulin. Among these, only one was common between the proteomes of renal failure and nephrotic syndrome patients. Among the limited proteins found in microalbuminuria patients, three were common with the proteome of nephrotic syndrome. Overall profiles were, however, quite different. Our study showed that urinary proteomes of different renal conditions were different and emphasized the potential of urinary proteome analysis to augment existing tools in the diagnosis of renal disorders. Biochemistry studies - General CC 10060 Biochemistry studies - Proteins, peptides and amino acids Pathology - Diagnostic 12504 Metabolism - Metabolic disorders 13020 Urinary system - Physiology and biochemistry 15504 Urinary system - Pathology 15506 IT Major Concepts Biochemistry and Molecular Biophysics; Nephrology (Human Medicine, Medical Sciences) TT Parts, Structures, & Systems of Organisms urine: excretory system IT Diseases kidney failure: urologic disease Kidney Failure (MeSH) ΙT Diseases microalbuminuria: metabolic disease, urologic disease Albuminuria (MeSH) ΙT

nephrotic syndrome: urologic disease

renal disorders: urologic disease

Nephrotic Syndrome (MeSH)

Chemicals & Biochemicals

TΤ

TΨ

Diseases

ANSWER 1 OF 17 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN AN 2002:566278 BIOSIS PREV200200566278 DN ΤI Proteomics of renal disorders: Urinary proteome analysis by two-dimensional gel electrophoresis and MALDI-TOF mass spectrometry. ΑU Kumar, Yadunanda; Uppuluri, Nageshwar Rao Venkata; Babu, Kishore; Phadke, Kishore; Kumar, Prasanna; Ballal, Sudarshan; Tatu, Utpal [Reprint author] CS Department of Biochemistry, Indian Institute of Science, Bangalore, 560 012, India tatu@biochem.iisc.ernet.in ·SO Current Science (Bangalore), (25 March, 2002) Vol. 82, No. 6, pp. 655-663. print. CODEN: CUSCAM. ISSN: 0011-3891. DTArticle LA English EDEntered STN: 7 Nov 2002 Last Updated on STN: 7 Nov 2002 The proteomes of urinary samples from patients with different renal conditions were analysed by two-dimensional electrophoresis and MALDI-TOF technology. Samples from three different renal conditions, namely kidney failure, nephrotic syndrome and microalbuminuria, were included in the analysis. Apart from the presence of albumin, the profiles of protein spots found in these urine samples were quite distinct. While kidney failure patients showed predominantly low molecular weight proteins, the nephrotic syndrome patients showed an abundance of relatively high molecular weight proteins clustering in the acidic range of the 2-D gels. Two different protein spots from kidney failure patients, four from nephrotic syndrome patients and three from micro-albuminuria patients were identified by in-gel protease digestions and analysis of resulting peptides by MALDI-TOF. The proteins identified were albumin, alpha-1-antitrypsin, alpha-1-acid glycoprotein 2, Zn-alpha-2-glycoprotein and alpha-1-microglobulin. Among these, only one was common between the proteomes of renal failure and nephrotic syndrome patients. Among the limited proteins found in microalbuminuria patients, three were common with the proteome of nephrotic syndrome. Overall profiles were, however, quite different. Our study showed that urinary proteomes of different renal conditions were different and emphasized the potential of urinary proteome analysis to augment existing tools in the diagnosis of renal disorders. CC Biochemistry studies - General 10060 Biochemistry studies - Proteins, peptides and amino acids Pathology - Diagnostic 12504 Metabolism - Metabolic disorders 13020 Urinary system - Physiology and biochemistry 15504 Urinary system - Pathology 15506 IT Major Concepts Biochemistry and Molecular Biophysics; Nephrology (Human Medicine, Medical Sciences) IT Parts, Structures, & Systems of Organisms urine: excretory system ITDiseases kidney failure: urologic disease Kidney Failure (MeSH) ΙT Diseases microalbuminuria: metabolic disease, urologic disease Albuminuria (MeSH) ΙT

nephrotic syndrome: urologic disease

renal disorders: urologic disease

Nephrotic Syndrome (MeSH)

Chemicals & Biochemicals

ΙT

·IT

albumin; alpha-1-acid glycoprotein 2; alpha-1-antitrypsin; alpha-1-microglobulin; high molecular weight proteins; low molecular weight proteins; zinc-alpha-2-glycoprotein

IT Methods & Equipment

matrix-assisted laser desorption ionization-time of flight mass spectrometry [MALDI-TOF mass spectrometry]: identification method; two-dimensional gel electrophoresis: molecular method, polyacrylamide gel electrophoresis; urinary proteome analysis: diagnostic method, molecular method

IT Miscellaneous Descriptors

proteomics

ORGN Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

human: patient

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates

albumin; alpha-1-acid glycoprotein 2; alpha-1-antitrypsin; alpha-1-microglobulin; high molecular weight proteins; low molecular weight proteins; zinc-alpha-2-glycoprotein

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